Audet 09/870087 Applicant

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2 E3-4 ه منسینی داند

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=> b hcap FILE 'HCAPLUS' ENTERED AT 14:03:38 ON 18 NOV 2004
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FILE COVERS 1907 - 18 Nov 2004 VOL 141 ISS 21 FILE LAST UPDATED: 17 Nov 2004 (20041117/ED)

This file contains CAS Registry Numbers for easy and accurate substance identification.

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- ANSWER 1 OF 2 HCAPLUS COPYRIGHT 2004 ACS on STN 1.1
- 2002:555963 HCAPLUS AN
- DN 137:114538
- Entered STN: 26 Jul 2002
- Ionic molecular conjugates of N-acylated derivatives of poly(2-amino-2-deoxy-D-glucose) and polypeptides
- Shalaby, Shalaby W.; Jackson, Steven A.; Ignatious, Francis X.; Moreau, Jacques-Pierre; Russell, Ruth M.
- PA
- U.S. Pat. Appl. Publ., 14 pp., Cont.-in-part of U.S. Ser. No. 929,363. SO CODEN: USXXCO
- DT Patent
- LA English
- ICM A61K009-00 IC
- NCL 424400000
- 63-6 (Pharmaceuticals)

Section cross-reference(s): 34

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	US 5665702	A	19970909	US 1995-468947	19950606
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Search done by Noble Jarrell

Page 1

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     WO 1999-US23406
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                         A61K009-00.
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                  ECLA
                         A61K038/31; C08B037/00M3B2
                 ECLA
                         A61K038/31; A61K047/48K8; C08B037/00M3B2; C08L005/08
 US 2003092800
   A copolymer comprising an N-acylated derivative, and a composition comprising said
     copolymer and a polypeptide, said polypeptide comprising at least one
     effective ionogenic amine, wherein at least 50 %, by weight, of said
     polypeptide present in said composition is ionically bound to said polymer.
     Conjugates were prepared from chitosan derivs. and a somatostatin
     polypeptide analog Somatuline.
     peptide acyl glucosamine polymer deriv conjugate; chitosan peptide
ST
     conjugate drug delivery
     Peptides, biological studies
IΤ
     RL: THU (Therapeutic use); BIOL (Biological study); USES (Uses)
         (conjugates; oral pharmaceutical dosage forms for pulsatile delivery of
        an antiarrhythmic agent)
IT
     Drug delivery systems
         (oral pharmaceutical dosage forms for pulsatile delivery of an
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     9012-76-4, Chitosan 9012-76-4D, Chitosan, N-succinylated
IT
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     108-30-5DP, Succinic anhydride, reaction products with depolymd. chitosan 108-55-4DP, Glutaric anhydride, reaction products with depolymd. chitosan
     123-62-6DP, Propionic anhydride, reaction products with depolymd. chitosan
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     57982-77-1DP, conjugates 64717-45-9DP, conjugates 65807-02-5DP, conjugates 66866-63-5DP, conjugates 76712-82-8DP, conjugates 78115-75-0DP, conjugates 127984-74-1DP, Somatuline, conjugates with acyl
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     conjugates 215945-52-1DP, conjugates
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         (oral pharmaceutical dosage forms for pulsatile delivery of an
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     9002-64-6, Parathyroid hormone
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L1
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AN
DN
     132:284253
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ED

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Entered STN: 21 Apr 2000
     Ionic molecular conjugates of N-acylated derivatives of
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     poly(2-amino-2-deoxy-D-glucose) and polypeptides
IN
     Shalaby, Shalaby W.; Jackson, Steven A.; Ignatious, Francis X.; Moreau,
     Jacques-Pierre; Russell, Ruth M.
     Societe De Conseils De Recherches Et D'applications Scientifiques S.A.,
PΔ
     Fr.
so
     PCT Int. Appl., 34 pp.
     CODEN: PIXXD2
     Patent
DT
     English
LA
IC
     ICM A61K047-36
     ICS A61K038-00; C08L005-08; C08B037-08
CC
     63-6 (Pharmaceuticals)
     Section cross-reference(s): 2, 33, 34
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 WO 2000021567
                  ICM
                         A61K047-36
                         A61K038-00; C08L005-08; C08B037-08
                  ICS
                         A61K038/31; C08B037/00M3B2
 US 2002098206
                 ECLA
     A copolymer comprises an N-acylated derivative, and a composition comprising said
     copolymer and a polypeptide, said polypeptide comprising at least one
     effective ionogenic amine, wherein at least 50 percent, by weight, of said
     polypeptide present in said composition is ionically bound to said polymer.
     Chitosan was depolymd., succinylated, , acetylated, and conjugated to the
     somatostatin peptide analog Somatuline.
ST
     aminodeoxyglucose polymer peptide conjugate
     Drug delivery systems
         (ionic mol. conjugates of N-acylated derivs. of poly(2-amino-2-deoxy-D-
         glucose) and polypeptides)
IT
     127984-74-1DP, Somatuline, conjugates with poly(N-acyl-D-glucosamine)s
     RL: SPN (Synthetic preparation); THU (Therapeutic use); BIOL (Biological study); PREP (Preparation); USES (Uses)
         (ionic mol. conjugates of N-acylated derivs. of poly(2-amino-2-deoxy-D-
        glucose) and polypeptides)
     108-30-5D, Succinic anhydride, reaction products with depolymd.chitosan,
     conjugates with peptides
                                  108-55-4D, Glutaric anhydride, reaction
     products with depolymd.chitosan, conjugates with peptides 123-62-6D,
     Propionic anhydride, reaction products with depolymd.chitosan, conjugates
                      9012-76-4D, Chitosan, depolymd., acyl derivs., conjugates
     with peptides
     with peptides 35110-26-0D, D-Glucose, 2-amino-2-deoxy-, homopolymer, N-acyl derivs., conjugates with peptides 38234-21-8D, Fertirelin,
     conjugates with poly (N-acyl-D-glucosamine)s 53714-56-0D, Leuprorelin,
     conjugates with poly(N-acyl-D-glucosamine)s
                                                      57773-63-4D, Tryptorelin,
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                                                       57773-65-6D, Deslorelin,
     conjugates with poly(N-acyl-D-glucosamine)s
                                                       57982-77-1D, Buserelin,
     conjugates with poly(N-acyl-D-glucosamine)s
                                                       65807-02-5D, Goserelin,
                                                       66866-63-5D, Lutrelin,
     conjugates with poly(N-acyl-D-glucosamine)s
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                                                       76712-82-8D, Histrelin,
                                                       76932-56-4D, Nafarelin,
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     poly (N-acyl-D-glucosamine) s
     RL: THU (Therapeutic use); BIOL (Biological study); USES (Uses)
        (ionic mol. conjugates of N-acylated derivs. of poly(2-amino-2-deoxy-D-
        glucose) and polypeptides)
RE.CNT 5
              THERE ARE 5 CITED REFERENCES AVAILABLE FOR THIS RECORD
RE
(1) Biomeasure Inc; WO 9504752 A 1995 HCAPLUS
(2) Kent, J; US 4675189 A 1987 HCAPLUS
(3) McNeil Ppc Inc; EP 0643963 A 1995 HCAPLUS
 (4) Shalaby, S; WO 9639160 A 1996 HCAPLUS
(5) Song, Y; JOURNAL OF CONTROLLED RELEASE V42(1), P93
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TSCA INFORMATION NOW CURRENT THROUGH MAY 21, 2004
  Please note that search-term pricing does apply when
  conducting SmartSELECT searches.
Crossover limits have been increased. See HELP CROSSOVER for details.
Experimental and calculated property data are now available. For more
information enter HELP PROP at an arrow prompt in the file or refer
to the file summary sheet on the web at:
http://www.cas.org/ONLINE/DBSS/registryss.html
--> d sqide 13 tot
L3
     ANSWER 1 OF 26 REGISTRY COPYRIGHT 2004 ACS on STN
     215945-52-1 REGISTRY
RN
     L-Threoninamide, N-[[4-(2-hydroxyethyl)-1-piperazinyl]acetyl]-D-
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OTHER NAMES:
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NTE modified (modifications unspecified)
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Sequence Patent
Source Reference
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Not Given US6004928
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RL.P Roles from patents: BIOL (Biological study); USES (Uses)

RLD.P Roles for non-specific derivatives from patents: BIOL (Biological study); PREP (Preparation); USES (Uses)

Absolute stereochemistry.

PAGE 1-B

\_NH2

PAGE 2-A

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- RN
- CN L-Threoninamide, N-[[2-[4-(2-hydroxyethyl)-1-piperazinyl]ethyl]sulfonyl]-Dphenylalanyl-L-phenylalanyl-L-phenylalanyl-D-tryptophyl-L-lysyl-L-threonyl-L-phenylalanyl- (9CI) (CA INDEX NAME)
  - PROTEIN SEQUENCE; STEREOSEARCH

FS SQL

modified

NIE MOUILIEU				
type	location		description	
terminal mod. modification	Thr-8 Phe-1	- -	C-terminal amide undetermined modification	

SEQ 1 FFFWKTFT

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C69 H91 N13 O13 S MF

SR CA

LC STN Files: CA, CAPLUS, TOXCENTER, USPAT2, USPATFULL

DT.CA CAplus document type: Patent

Roles from patents: BIOL (Biological study); USES (Uses)

RLD.P Roles for non-specific derivatives from patents: BIOL (Biological study); PREP (Preparation); USES (Uses)

Absolute stereochemistry.

PAGE 1-B

PAGE 2-A

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ANSWER 3 OF 26 REGISTRY COPYRIGHT 2004 ACS on STN L3 RN

204388-14-7 REGISTRY

L-Threoninamide, N-[[4-(2-hydroxyethyl)-1-piperazinyl]acetyl]-D-CN phenylalanyl-L-cysteinyl-L-tyrosyl-D-tryptophyl-L-lysyl-(2S)-2aminobutanoyl-L-cysteinyl- (9CI) (CA INDEX NAME)

PROTEIN SEQUENCE; STEREOSEARCH FS

SQL

modified (modifications unspecified)

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stereo	Tyr-5	-	D		

#### SEQ 1 GFCYWKXCT

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Absolute stereochemistry.

PAGE 1-A

PAGE 1-B

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- 5 REFERENCES IN FILE CA (1907 TO DATE)
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- 5 REFERENCES IN FILE CAPLUS (1907 TO DATE)
- L3 ANSWER 4 OF 26 REGISTRY COPYRIGHT 2004 ACS on STN
- RN 204388-13-6 REGISTRY
- CN L-Threoninamide, N-[[2-[4-(2-hydroxyethyl)-1-piperazinyl]ethyl]sulfonyl]-Dphenylalanyl-L-cysteinyl-L-tyrosyl-D-tryptophyl-L-lysyl-(2S)-2aminobutanoyl-L-cysteinyl- (9CI) (CA INDEX NAME)
- FS PROTEIN SEQUENCE; STEREOSEARCH

ОН

- SQL 8
- NTE modified

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uncommon	Abu-6	-	
modification	Phe-1	-	

#### SEQ 1 FCYWKXCT

\*\*RELATED SEQUENCES AVAILABLE WITH SEQLINK\*\*

C57 H83 N13 O13 S3 MF

SR CA

LC STN Files: CA, CAPLUS, TOXCENTER, USPAT2, USPATFULL
DT.CA CAplus document type: Patent
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RLD.P Roles for non-specific derivatives from patents: BIOL (Biological study); PREP (Preparation); USES (Uses)

Absolute stereochemistry.

PAGE 1-A

PAGE 1-B

- 6 REFERENCES IN FILE CA (1907 TO DATE)
- 2 REFERENCES TO NON-SPECIFIC DERIVATIVES IN FILE CA
- 6 REFERENCES IN FILE CAPLUS (1907 TO DATE)
- L3 ANSWER 5 OF 26 REGISTRY COPYRIGHT 2004 ACS on STN
- 148440-40-8 REGISTRY RN
- CN L-Threoninamide, 3-(1-naphthalenyl)-D-alanyl-L-cysteinyl-L-tyrosyl-Dtryptophyl-L-lysyl-L-valyl-L-cysteinyl-, cyclic (2.fwdarw.7)-disulfide (9CI) (CA INDEX NAME)

OTHER CA INDEX NAMES:

1,2-Dithia-5,8,11,14,17-pentaazacycloeicosane, cyclic peptide deriv. PROTEIN SEQUENCE; STEREOSEARCH CN

FS

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SQL 8
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terminal mod.
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bridge
modification
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SEO
**RELATED SEQUENCES AVAILABLE WITH SEQLINK**
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MF
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COM CI

SR CA

LC STN Files: CA, CAPLUS, TOXCENTER, USPAT2, USPATFULL DT.CA CAplus document type: Journal; Patent

Roles from patents: BIOL (Biological study); USES (Uses) RL.P

RLD.P Roles for non-specific derivatives from patents: BIOL (Biological

study); PREP (Preparation); USES (Uses) RL.NP Roles from non-patents: RACT (Reactant or reagent)

- 4 REFERENCES IN FILE CA (1907 TO DATE)
- 2 REFERENCES TO NON-SPECIFIC DERIVATIVES IN FILE CA
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L3 ANSWER 6 OF 26 REGISTRY COPYRIGHT 2004 ACS on STN

132609-33-7 REGISTRY RN

L-Threoninamide, 3-(1-naphthalenyl)-D-alanyl-L-cysteinyl-L-tyrosyl-D-CN tryptophyl-L-lysyl-L-valyl-L-cysteinyl- (9CI) (CA INDEX NAME)

OTHER NAMES:

CN Lantreotide

PROTEIN SEQUENCE; STEREOSEARCH FS

SQL 8

NTE modified

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type	loc	ation	description	
terminal mod. modification	Thr-8 Ala-1	- -	C-terminal amide 1-naphthalenyl<1-Naph>	

#### SEQ 1 ACYWKVCT

\*\*RELATED SEQUENCES AVAILABLE WITH SEQLINK\*\*

MF C54 H71 N11 O10 S2

SR CA

LC STN Files: BIOSIS, CA, CAPLUS, TOXCENTER, USPATZ, USPATFULL DT.CA CAplus document type: Journal; Patent RL.P Roles from patents: BIOL (Biological study)

RLD.P Roles for non-specific derivatives from patents: BIOL (Biological study); PREP (Preparation); USES (Uses)

RL.NP Roles from non-patents: ANST (Analytical study); BIOL (Biological study)

Absolute stereochemistry.

PAGE 1-A

PAGE 2-A

- 5 REFERENCES IN FILE CA (1907 TO DATE) 2 REFERENCES TO NON-SPECIFIC DERIVATIVES IN FILE CA
- 5 REFERENCES IN FILE CAPLUS (1907 TO DATE)
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OTHER CA INDEX NAMES:

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- CN 2: PN: WO0006185 PAGE: 8 claimed protein
- CN BIM 23014C
- CN Lanreotide acetate Somatulina
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- PROTEIN SEQUENCE; STEREOSEARCH FS

SQL 8

NTE modified

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modification	-	-	
modification	Ala-1	-	

PATENT ANNOTATIONS (PNTE):

Sequence | Patent Source Reference  Not Given WO2000006185 claimed PAGE 8

SEQ 1 ACYWKVCT

\*\*RELATED SEQUENCES AVAILABLE WITH SEQLINK\*\*

MF C54 H69 N11 O10 S2 . x C2 H4 O2

SR CA

LC STN Files: BIOBUSINESS, BIOSIS, CA, CAPLUS, CIN, DDFU, DRUGU,
IMSCOSEARCH, IMSPATENTS, IMSRESEARCH, IPA, MRCK\*, PROMT, PROUSDDR,
TOXCENTER, USAN, USPAT2, USPATFULL

(\*File contains numerically searchable property data)

DT.CA CAplus document type: Journal; Patent

RL.P Roles from patents: BIOL (Biological study); PROC (Process); RACT (Reactant or reagent); USES (Uses)

RLD.P Roles for non-specific derivatives from patents: BIOL (Biological study); PREP (Preparation); RACT (Reactant or reagent); USES (Uses)

RL.NP Roles from non-patents: BIOL (Biological study); PROC (Process); PRP (Properties); USES (Uses)

CM 1

CRN 108736-35-2 CMF C54 H69 N11 O10 S2

CM 2

CRN 64-19-7 CMF C2 H4 O2

- 47 REFERENCES IN FILE CA (1907 TO DATE)
- 3 REFERENCES TO NON-SPECIFIC DERIVATIVES IN FILE CA
- 47 REFERENCES IN FILE CAPLUS (1907 TO DATE)

L3 ANSWER 8 OF 26 REGISTRY COPYRIGHT 2004 ACS on STN

RN 113294-82-9 REGISTRY

CN L-Threoninamide, 3-(2-naphthalenyl)-D-alanyl-L-cysteinyl-L-tyrosyl-D-tryptophyl-L-lysyl-L-valyl-L-cysteinyl- (9CI) (CA INDEX NAME)

FS PROTEIN SEQUENCE; STEREOSEARCH

SQL 8

NTE modified

type ----- location ---- description

terminal mod. Thr-8 - C-terminal amide
modification Ala-1 - 2-naphthalenyl<2-Naph>

•

-----

## SEQ 1 ACYWKVCT

\*\*RELATED SEQUENCES AVAILABLE WITH SEQLINK\*\*

MF C54 H71 N11 O10 S2

SR CA

LC STN Files: BIOTECHNO, CA, CANCERLIT, CAPLUS, EMBASE, MEDLINE, TOXCENTER, USPATFULL

DT.CA CAplus document type: Journal; Patent

RL.P Roles from patents: BIOL (Biological study); PREP (Preparation); USES (Uses)

RLD.P Roles for non-specific derivatives from patents: BIOL (Biological study); USES (Uses)

RL.NP Roles from non-patents: BIOL (Biological study); PREP (Preparation); RACT (Reactant or reagent); USES (Uses)

## Absolute stereochemistry.

PAGE 1-A

PAGE 1-B

- 10 REFERENCES IN FILE CA (1907 TO DATE)
- 3 REFERENCES TO NON-SPECIFIC DERIVATIVES IN FILE CA
- 10 REFERENCES IN FILE CAPLUS (1907 TO DATE)

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L3
    ANSWER 9 OF 26 REGISTRY COPYRIGHT 2004 ACS on STN
    78115-75-0 REGISTRY
RN
    Luteinizing hormone-releasing factor (swine), 6-[3-(1-naphthalenyl)-D-
CN
     alanine] - (9CI) (CA INDEX NAME)
OTHER CA INDEX NAMES:
    Luteinizing hormone-releasing factor (pig), 6-[3-(1-naphthalenyl)-D-
CN
     alanine]-
     PROTEIN SEQUENCE; STEREOSEARCH
FS
SQL 10
NTE modified
                ----- location -----
type
                                             description
terminal mod.
                Gly-10
                                          C-terminal amide
uncommon
                Glp-1
modification
                Ala-6
                                          1-naphthalenyl<1-Naph>
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1 XHWSYALRPG SEO

\*\*RELATED SEQUENCES AVAILABLE WITH SEQLINK\*\*

MF C66 H83 N17 O13

CI COM

LC STN Files: CA, CAPLUS, USPAT2, USPATFULL DT.CA CAplus document type: Conference; Journal; Patent

Roles from patents: BIOL (Biological study); PREP (Preparation); USES RL.P (Uses)

Roles for non-specific derivatives from patents: BIOL (Biological RLD.P study); PREP (Preparation); USES (Uses)

RL.NP Roles from non-patents: BIOL (Biological study); PREP (Preparation)

Absolute stereochemistry. Rotation (-).

PAGE 1-A

PAGE 1-B

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      HN 
              NH<sub>2</sub>
                6 REFERENCES IN FILE CA (1907 TO DATE)
                1 REFERENCES TO NON-SPECIFIC DERIVATIVES IN FILE CA
                6 REFERENCES IN FILE CAPLUS (1907 TO DATE)
     ANSWER 10 OF 26 REGISTRY COPYRIGHT 2004 ACS on STN
L3
     76932-56-4 REGISTRY
RN
     Luteinizing hormone-releasing factor (swine), 6-[3-(2-naphthalenyl)-D-
CN
     alanine] - (9CI) (CA INDEX NAME)
OTHER CA INDEX NAMES:
     Luteinizing hormone-releasing factor (pig), 6-[3-(2-naphthalenyl)-D-
     alaninel-
OTHER NAMES:
CN
    Nafarelin
     Nafareline
CN
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NTE modified
          ----- location ----- description
type
  .
          Gly-10
                            C-terminal amide
terminal mod.
uncommon
           Glp-1
                           2-naphthalenyl<2-Naph>
modification
          Ala-6
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#### SEO 1 XHWSYALRPG

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**RELATED SEQUENCES AVAILABLE WITH SEQLINK**
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[6-D-(2-naphthyl)-alanine]LH-RH

PROTEIN SEQUENCE; STEREOSEARCH

80458-30-6

MF C66 H83 N17 O13

COM CI

CN

CN

FS SQL 10

NAG

TN Files: ADISNEWS, AGRICOLA, ANABSTR, BIOBUSINESS, BIOSIS, BIOTECHNO, CA, CANCERLIT, CAPLUS, CBNB, CHEMCATS, CIN, DDFU, DRUGU, EMBASE, LC STN Files: IMSDRUGNEWS, IMSPATENTS, IMSRESEARCH, IPA, MEDLINE, MRCK\*, PHAR, PROMT, PROUSDDR, RTECS\*, TOXCENTER, USAN, USPAT2, USPATFULL, VETU

(\*File contains numerically searchable property data) WHO

Other Sources: DT.CA CAplus document type: Conference; Journal; Patent

Roles from patents: BIOL (Biological study); PREP (Preparation); PROC RL.P (Process); PRP (Properties); RACT (Reactant or reagent); USES (Uses)

Roles for non-specific derivatives from patents: BIOL (Biological study); PREP (Preparation); PRP (Properties); USES (Uses)

Roles from non-patents: ANST (Analytical study); BIOL (Biological RL.NP study); PREP (Preparation); PROC (Process); PRP (Properties); USES

RLD.NP Roles for non-specific derivatives from non-patents: BIOL (Biological study)

Absolute stereochemistry.

PAGE 1-A

PAGE 1-B

228 REFERENCES IN FILE CA (1907 TO DATE)

7 REFERENCES TO NON-SPECIFIC DERIVATIVES IN FILE CA

229 REFERENCES IN FILE CAPLUS (1907 TO DATE)

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ANSWER 11 OF 26 REGISTRY COPYRIGHT 2004 ACS on STN
L3
     76712-82-8 REGISTRY
RN
    1-9-Luteinizing hormone-releasing factor (swine), 6-[1-(phenylmethyl)-D-
    histidine] -9-(N-ethyl-L-prolinamide) - (9CI) (CA INDEX NAME)
OTHER CA INDEX NAMES:
    Luteinizing hormone-releasing factor (pig), 6-[1-(phenylmethyl)-D-
    histidine] -9- (N-ethyl-L-prolinamide) -10-deglycinamide-
OTHER NAMES:
CN
    Histrelin
CN
    ORF 17070
CN
    RWJ 17070
    PROTEIN SEQUENCE; STEREOSEARCH
FS
SOL
    modified (modifications unspecified)
NTE
                ----- location -----
                                              description
 type
          Glp-1
uncommon
modification
                His-6
                                         phenylmethyl<Bzl>
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1 XHWSYHLRP SEQ

\*\*RELATED SEQUENCES AVAILABLE WITH SEQLINK\*\*

DR 97708-83-3, 102989-36-6

C66 H86 N18 O12 MF

COM CI

STN Files: ADISNEWS, BEILSTEIN\*, BIOBUSINESS, BIOSIS, BIOTECHNO, CA, LC CANCERLIT, CAPLUS, CHEMCATS, CIN, DDFU, DRUGU, EMBASE, IMSDRUGNEWS, IMSPATENTS, IMSRESEARCH, IPA, MEDLINE, MRCK\*, PHAR, PROMT, PROUSDDR, RTECS\*, TOXCENTER, USAN, USPAT2, USPATFULL, VETU

(\*File contains numerically searchable property data)

Other Sources: WHO

DT.CA CAplus document type: Journal; Patent

Roles from patents: BIOL (Biological study); PROC (Process); RACT RL.P (Reactant or reagent); USES (Uses)

RLD.P Roles for non-specific derivatives from patents: BIOL (Biological study); PREP (Preparation); USES (Uses)

RL.NP Roles from non-patents: BIOL (Biological study); PREP (Preparation); PROC (Process); PRP (Properties); USES (Uses)

Absolute stereochemistry.

OH

PAGE 1-B

PAGE 1-A

119 REFERENCES IN FILE CA (1907 TO DATE)

5 REFERENCES TO NON-SPECIFIC DERIVATIVES IN FILE CA 120 REFERENCES IN FILE CAPLUS (1907 TO DATE)

ANSWER 12 OF 26 REGISTRY COPYRIGHT 2004 ACS on STN L3

66866-63-5 REGISTRY

RN

1-9-Luteinizing hormone-releasing factor (swine), 6-D-tryptophan-7-(N-methyl-L-leucine)-9-(N-ethyl-L-prolinamide)- (9CI) (CA INDEX NAME) CN

Search done by Noble Jarrell

```
OTHER CA INDEX NAMES:
    Luteinizing hormone-releasing factor (pig), 6-D-tryptophan-7-(N-methyl-L-
     leucine) -9-(N-ethyl-L-prolinamide) -10-deglycinamide-
OTHER NAMES:
CN
    Lutrelin
CN
    Wy 40972
    Wyeth 40972
CN
     [D-Trp6-N-methyl-Leu7-des-Gly10-Pro9-NH]-LH-RH ethylamide
CN
    PROTEIN SEQUENCE; STEREOSEARCH
FS
SQL
NTE modified (modifications unspecified)
_____
type
                ----- location -----
                                              description
                Glp-1
uncommon
modification
                Leu-7
                                         methyl<Me>
        1 XHWSYWLRP
SEO
**RELATED SEQUENCES AVAILABLE WITH SEQLINK**
   102586-12-9, 67910-57-0
DR
    C65 H85 N17 O12
MF
CI
    COM
                BIOSIS, BIOTECHNO, CA, CANCERLIT, CAPLUS, DDFU, DRUGU,
LC
     STN Files:
      EMBASE, IFICDB, IFIPAT, IFIUDB, MEDLINE, PHAR, PROUSDDR, RTECS*, TOXCENTER, USAN, USPAT2, USPATFULL
         (*File contains numerically searchable property data)
     Other Sources:
                    WHO
DT.CA CAplus document type: Journal; Patent
       Roles from patents: BIOL (Biological study); PREP (Preparation); PROC
RL.P
       (Process); USES (Uses)
RLD.P Roles for non-specific derivatives from patents: BIOL (Biological
       study); PREP (Preparation); USES (Uses)
      Roles from non-patents: BIOL (Biological study); PREP (Preparation);
RL.NP
       PROC (Process)
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Absolute stereochemistry.

PAGE 1-B

PAGE 2-A

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NH2
(CH2) 3
                NHET
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79 REFERENCES IN FILE CA (1907 TO DATE)
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- 3 REFERENCES TO NON-SPECIFIC DERIVATIVES IN FILE CA
- 79 REFERENCES IN FILE CAPLUS (1907 TO DATE)

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ANSWER 13 OF 26 REGISTRY COPYRIGHT 2004 ACS on STN
L3
    65807-02-5 REGISTRY
ВM
    1-9-Luteinizing hormone-releasing factor (swine), 6-[0-(1,1-dimethylethyl)-
CN
    D-serine]-, 2-(aminocarbonyl)hydrazide (9CI) (CA INDEX NAME)
OTHER CA INDEX NAMES:
    Luteinizing hormone-releasing factor (pig), 6-[0-(1,1-dimethylethyl)-D-
    serine]-10-deglycinamide-, 2-(aminocarbonyl)hydrazide
OTHER NAMES:
    Decapeptide I
CN
    Goserelin
CN
CN
    ICI 118630
CN
    Zoladex
    PROTEIN SEQUENCE; STEREOSEARCH
FS
SOL 9
NTE modified (modifications unspecified)
                ----- location -----
                                            description
type
               -----
uncommon Glp-1
modification
                Ser-6
                                         1,1-dimethylethyl<t-Bu>
        1 XHWSYSLRP
SEO
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**RELATED SEQUENCES AVAILABLE WITH SEQLINK**
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70280-59-0 DR

MF C59 H84 N18 O14

CI COM

ADISINSIGHT, ADISNEWS, AGRICOLA, ANABSTR, BIOBUSINESS, BIOSIS, BIOTECHNO, CA, CANCERLIT, CAPLUS, CASREACT, CBNB, CEN, CHEMCATS, CHEMLIST, CIN, DDFU, DIOGENES, DRUGU, EMBASE, IFICDB, IFIPAT, IFIUDB, IMSCOSEARCH, IMSDRUGNEWS, IMSPATENTS, IMSRESEARCH, IPA, MEDLINE, MRCK\*, PHAR, PROMT, PROUSDDR, PS, RTECS\*, TOXCENTER, USAN, USPAT2, USPATFULL, VETU

(\*File contains numerically searchable property data)

Other Sources: WHO

DT.CA CAplus document type: Conference; Journal; Patent

Roles from patents: BIOL (Biological study); PREP (Preparation); PROC RL.P (Process); PRP (Properties); RACT (Reactant or reagent); USES (Uses)

Roles for non-specific derivatives from patents: BIOL (Biological RLD.P study); PREP (Preparation); USES (Uses)

Roles from non-patents: ANST (Analytical study); BIOL (Biological study); PREP (Preparation); PROC (Process); PRP (Properties); USES (Uses)

RLD.NP Roles for non-specific derivatives from non-patents: PRP (Properties)

Absolute stereochemistry.

PAGE 1-A

PAGE 1-B

489 REFERENCES IN FILE CA (1907 TO DATE)

10 REFERENCES TO NON-SPECIFIC DERIVATIVES IN FILE CA

492 REFERENCES IN FILE CAPLUS (1907 TO DATE)

L3 ANSWER 14 OF 26 REGISTRY COPYRIGHT 2004 ACS on STN

RN 64717-45-9 REGISTRY

CN 1-9-Luteinizing hormone-releasing factor (swine), 6-L-tryptophan-9-(Nethyl-L-prolinamide)- (9CI) (CA INDEX NAME)

OTHER CA INDEX NAMES:

CN Luteinizing hormone-releasing factor (pig), 6-L-tryptophan-9-(N-ethyl-L-prolinamide)-10-deglycinamide-

FS PROTEIN SEQUENCE; STEREOSEARCH

SQL 9

NTE modified (modifications unspecified)

type ----- location ----- description
uncommon Glp-1 - -

SEQ 1 XHWSYWLRP

\*\*RELATED SEQUENCES AVAILABLE WITH SEQLINK\*\*

MF C64 H83 N17 O12

CI COM

LC STN Files: CA, CAPLUS, IFICDB, IFIPAT, IFIUDB, TOXCENTER, USPAT2, USPATFULL

DT.CA CAplus document type: Journal; Patent

RL.P Roles from patents: PREP (Preparation)

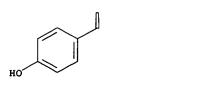
RLD.P Roles for non-specific derivatives from patents: BIOL (Biological study); PREP (Preparation); USES (Uses)

RL.NP Roles from non-patents: BIOL (Biological study) Absolute stereochemistry.

PAGE 1-A

PAGE 1-B

PAGE 2-A



- 5 REFERENCES IN FILE CA (1907 TO DATE) 2 REFERENCES TO NON-SPECIFIC DERIVATIVES IN FILE CA
- 5 REFERENCES IN FILE CAPLUS (1907 TO DATE)
- ANSWER 15 OF 26 REGISTRY COPYRIGHT 2004 ACS on STN 57982-77-1 REGISTRY L3
- RN
- 1-9-Luteinizing hormone-releasing factor (swine), 6-[0-(1,1-dimethylethyl)-D-serine] -9-(N-ethyl-L-prolinamide) - (9CI) (CA INDEX NAME)
  OTHER CA INDEX NAMES:

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Luteinizing hormone-releasing factor (pig), 6-[0-(1,1-dimethylethyl)-D-
CN
     serine] -9- (N-ethyl-L-prolinamide) -10-deglycinamide-
OTHER NAMES:
    1-9-(D-Ser(t-butyl))6-LH-releasing hormone ethylamide
CN
CN
    Buserelin
CN
    Etilamide
    HOE 766
CN
    HOE 766A
CN
CN
    ICI 123215
CN
    Receptal
CN
    Suprefact
     [D-Ser(tert-butyl)6,des-Gly-NH210]-LH-RH ethylamide [D-Ser6(t-Bu),de-Gly10-NH2]-LH-RH ethylamide
CN
CN
     PROTEIN SEQUENCE; STEREOSEARCH
SOL
NTE
    modified (modifications unspecified)
______
                ----- location ----- description
type
              Glp-1
uncommon
modification
                 Ser-6
                                            1,1-dimethylethyl<t-Bu>
         1 XHWSYSLRP
SEO
**RELATED SEQUENCES AVAILABLE WITH SEQLINK**
     476329-44-9, 121698-99-5, 102586-11-8, 104428-01-5, 111520-35-5,
DR
     70910-44-0
MF
    C60 H86 N16 O13
CI
     COM
    STN Files:
                 ADISNEWS, AGRICOLA, ANABSTR, BIOBUSINESS, BIOSIS, BIOTECHNO,
       CA, CANCERLIT, CAPLUS, CASREACT, CBNB, CHEMLIST, CIN, CSCHEM, DDFU,
       DRUGU, EMBASE, IFICDB, IFIPAT, IFIUDB, IMSCOSEARCH, IMSPATENTS, IPA, MEDLINE, MRCK*, PHAR, PROMT, PROUSDDR, PS, RTECS*, TOXCENTER, USAN,
       USPAT2, USPATFULL, VETU
         (*File contains numerically searchable property data)
     Other Sources: EINECS**, WHO
         (**Enter CHEMLIST File for up-to-date regulatory information)
       CAplus document type: Conference; Dissertation; Journal; Patent
       Roles from patents: BIOL (Biological study); PREP (Preparation); PROC
RL.P
       (Process); RACT (Reactant or reagent); USES (Uses)
RLD.P Roles for non-specific derivatives from patents: BIOL (Biological
       study); PREP (Preparation); USES (Uses)
RL.NP
      Roles from non-patents: ANST (Analytical study); BIOL (Biological
       study); PREP (Preparation); PROC (Process); PRP (Properties); RACT
       (Reactant or reagent); USES (Uses)
RLD.NP Roles for non-specific derivatives from non-patents: ANST (Analytical
       study); BIOL (Biological study); PREP (Preparation); PROC (Process)
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Absolute stereochemistry.

PAGE 1-B

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NHET NH2

L3 ANSWER RN 57773-6
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1041 REFERENCES IN FILE CA (1907 TO DATE)
               12 REFERENCES TO NON-SPECIFIC DERIVATIVES IN FILE CA
             1042 REFERENCES IN FILE CAPLUS (1907 TO DATE)
     ANSWER 16 OF 26 REGISTRY COPYRIGHT 2004 ACS on STN
    57773-65-6 REGISTRY
     1-9-Luteinizing hormone-releasing factor (swine), 6-D-tryptophan-9-(N-
CN
     ethyl-L-prolinamide) - (9CI) (CA INDEX NAME)
OTHER CA INDEX NAMES:
    Luteinizing hormone-releasing factor (pig), 6-D-tryptophan-9-(N-ethyl-L-
     prolinamide) -10-deglycinamide-
OTHER NAMES:
CN
    Bachem 9022
    D-Trp LHRH-PEA
CN
     D-Trp6-Pro9-N-ethylamide-LH-RH
CN
    Deslorelin
CN
CN
     H 4065
CN
     PTL 3001
CN
     Somagard
CN
     Somagorad
     [D-Trp6,des-Gly-NH210]-LH-RH ethylamide
CN
CN
     [D-Trp6, des-Gly10] -LH-RH ethylamide
     [D-Trp6, Pro9-NHEt] LH-RH
CN
     [de-Gly10, D-Trp6, Pro-NHEt] -LH-RH
CN
     [Des-Gly10[D-Trp6]-LH-RH ethylamide
CN
FS
     PROTEIN SEQUENCE; STEREOSEARCH
NTE modified (modifications unspecified)
                 ----- location -----
                                                 description
uncommon Glp-1
SEQ · 1 XHWSYWLRP
**RELATED SEQUENCES AVAILABLE WITH SEQLINK**
DR
     67190-19-6
ΜF
     C64 H83 N17 O12
CI
       TN Files: ADISINSIGHT, ADISNEWS, AGRICOLA, ANABSTR, BIOBUSINESS, BIOSIS, BIOTECHNO, CA, CANCERLIT, CAPLUS, CASREACT, CBNB, CHEMCATS, CIN,
LC
     STN Files:
       CSCHEM, DDFU, DRUGU, EMBASE, IFICDB, IFIPAT, IFIUDB, IMSDRUGNEWS
       IMSPATENTS, IMSRESEARCH, IPA, MEDLINE, MRCK*, PHAR, PROMT, PROUSDDR, PS,
       RTECS*, TOXCENTER, USAN, USPAT2, USPATFULL
          (*File contains numerically searchable property data)
     Other Sources: WHO
DT.CA Caplus document type: Conference; Dissertation; Journal; Patent; Report
       Roles from patents: BIOL (Biological study); PREP (Preparation); PROC
RL.P
        (Process); RACT (Reactant or reagent); USES (Uses)
RLD.P Roles for non-specific derivatives from patents: BIOL (Biological study); PREP (Preparation); PROC (Process); USES (Uses)
RL.NP Roles from non-patents: ANST (Analytical study); BIOL (Biological
       study); OCCU (Occurrence); PREP (Preparation); PROC (Process); PRP (Properties); RACT (Reactant or reagent); USES (Uses)
```

Absolute stereochemistry.

PAGE 1-A

PAGE 1-B



327 REFERENCES IN FILE CA (1907 TO DATE)
7 REFERENCES TO NON-SPECIFIC DERIVATIVES IN FILE CA 328 REFERENCES IN FILE CAPLUS (1907 TO DATE)

- ANSWER 17 OF 26 REGISTRY COPYRIGHT 2004 ACS on STN 57773-63-4 REGISTRY L3
- RN
- Luteinizing hormone-releasing factor (swine), 6-D-tryptophan- (9CI) (CA INDEX NAME)

OTHER CA INDEX NAMES:

- Luteinizing hormone-releasing factor (pig), 6-D-tryptophan-CN OTHER NAMES:
- 6-D-Tryptophan-LH-RH CN

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AY 25650
CN
CN
      CL 118532
     D-Tryptophan6-LH-RH
CN
     Triptorelin
CN
CN
     Triptoreline
CN
     Tryptorelin
     Wy 42422
Wy 42462
CN
CN
      [6-D-Tryptophan] luteinizing hormone-releasing hormone
CN
CN
      [D-Trp6]-GnRH
      PROTEIN SEQUENCE; STEREOSEARCH
FS
     10
SOL
NTE modified
                    ----- location ----- description
 type
terminal mod. Gly-10 - C-terminal amide
                   Glp-1
uncommon
SEO
          1 XHWSYWLRPG
**RELATED SEQUENCES AVAILABLE WITH SEQLINK**
MF
     C64 H82 N18 O13
CI
      COM
LC
      STN Files:
                     ADISINSIGHT, ADISNEWS, AGRICOLA, ANABSTR, BIOBUSINESS,
        BIOSIS, BIOTECHNO, CA, CANCERLIT, CAPLUS, CASREACT, CBNB, CHEMCATS, CIN, CSCHEM, DDFU, DIOGENES, DRUGU, EMBASE, IFICDB, IFIPAT, IFIUDB, IMSCOSEARCH, IMSDRUGNEWS, IMSPATENTS, IMSRESEARCH, IPA, MEDLINE, MRCK*,
        PHAR, PROMT, PROUSDDR, TOXCENTER, USAN, USPAT2, USPATFULL, VETU
           (*File contains numerically searchable property data)
      Other Sources: WHO
DT.CA CAplus document type: Conference; Journal; Patent RL.P Roles from patents: BIOL (Biological study); PREP (Preparation); PROC
        (Process); RACT (Reactant or reagent); USES (Uses)
RLD.P Roles for non-specific derivatives from patents: ANST (Analytical
        study); BIOL (Biological study); PREP (Preparation); PROC (Process); PRP
        (Properties); USES (Uses)
RL.NP Roles from non-patents: ANST (Analytical study); BIOL (Biological
        study); PREP (Preparation); PROC (Process); PRP (Properties); RACT (Reactant or reagent); USES (Uses)
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RLD.NP Roles for non-specific derivatives from non-patents: ANST (Analytical

Absolute stereochemistry. Rotation (-).

study); PROC (Process); PRP (Properties)

PAGE 1-B

PAGE 2-A

583 REFERENCES IN FILE CA (1907 TO DATE)

13 REFERENCES TO NON-SPECIFIC DERIVATIVES IN FILE CA

585 REFERENCES IN FILE CAPLUS (1907 TO DATE)

L3 ANSWER 18 OF 26 REGISTRY COPYRIGHT 2004 ACS on STN

RN 53714-56-0 REGISTRY

1-9-Luteinizing hormone-releasing factor (swine), 6-D-leucine-9-(N-ethyl-Lprolinamide) - (9CI) (CA INDEX NAME)

OTHER CA INDEX NAMES:

Luteinizing hormone-releasing factor (pig), 6-D-leucine-9-(N-ethyl-L-CN prolinamide) -10-deglycinamide-

OTHER NAMES:

(D-Leu6, des-Gly-NH210)-LH-RH ethylamide 1: PN: WO02087616 PAGE: 31 claimed protein CN

CN

CN

D-Leu6-des-Gly10-LH-releasing hormone ethylamide Des-Gly10-[D-Leu6]-LH-releasing hormone ethylamide CN

CN

CN Des-Gly10-[D-Leu6]LH-RH ethylamide

CNLeuprolide

CN Leuprorelin

CN Lupron SR

NSC 377526 CN

CNPGlu-His-Trp-Ser-Tyr-D-Leu-Leu-Arg-Pro-NHC2H5

FS PROTEIN SEQUENCE; STEREOSEARCH

SOL

modified (modifications unspecified) NTE

----- location ----type description Glp-1

uncommon

PATENT ANNOTATIONS (PNTE): Sequence | Patent Source Reference -----Not Given W02002087616

| claimed PAGE 31

SEO 1 XHWSYLLRP

\*\*RELATED SEQUENCES AVAILABLE WITH SEQLINK\*\*

DR 102586-10-7, 71873-71-7, 72648-87-4

MF C59 H84 N16 O12

CI COM

LC STN Files: ADISINSIGHT, ADISNEWS, AGRICOLA, ANABSTR, BEILSTEIN\*,
BIOBUSINESS, BIOSIS, BIOTECHNO, CA, CANCERLIT, CAPLUS, CBNB, CEN,
CHEMCATS, CIN, CSCHEM, DDFU, DIOGENES, DRUGU, EMBASE, HSDB\*, IFICDB,
IFIPAT, IFIUDB, IMSDRUGNEWS, IMSPATENTS, IMSRESEARCH, IPA, MEDLINE,
MRCK\*, PHAR, PROMT, PROUSDDR, PS, RTECS\*, TOXCENTER, USPAT2, USPATFULL,
VETU

(\*File contains numerically searchable property data)

Other Sources: WHO

DT.CA CAplus document type: Conference; Dissertation; Journal; Patent RL.P Roles from patents: BIOL (Biological study); PREP (Preparation); PROC (Process); PRP (Properties); RACT (Reactant or reagent); USES (Uses)

RLD.P Roles for non-specific derivatives from patents: ANST (Analytical study); BIOL (Biological study); PREP (Preparation); PRP (Properties); USES (Uses)

RL.NP Roles from non-patents: ANST (Analytical study); BIOL (Biological study); PREP (Preparation); PROC (Process); PRP (Properties); RACT (Reactant or reagent); USES (Uses)

RLD.NP Roles for non-specific derivatives from non-patents: ANST (Analytical study); PRP (Properties)

Absolute stereochemistry. Rotation (-).

PAGE 1-A

PAGE 1-B

702 REFERENCES IN FILE CA (1907 TO DATE)

17 REFERENCES TO NON-SPECIFIC DERIVATIVES IN FILE CA

706 REFERENCES IN FILE CAPLUS (1907 TO DATE)

```
ANSWER 19 OF 26 REGISTRY COPYRIGHT 2004 ACS on STN
L3
     51110-01-1 REGISTRY
RN
     Somatostatin (9CI) (CA INDEX NAME)
CN
OTHER NAMES:
CN
    Aminopan
CN
    AY 24910
    GH-RIF
CN
    Growth hormone release-inhibiting factor
CN
CN
    Growth hormone release-inhibiting hormone
CN
    Panhibin
CN
     Somatostatin-14
CN
     Somatotropin release-inhibiting factor
CN
     Somatotropin release-inhibiting hormone
CN
CN
     Somiaton
CN
     SRIF
CN
     SRIF 14
     56451-83-3, 52500-64-8, 53126-12-8
DR
MF
     Unspecified
CI
     MAN
                 ADISNEWS, AGRICOLA, ANABSTR, BIOBUSINESS, BIOSIS, BIOTECHNO,
     STN Files:
LC
       CA, CABA, CANCERLIT, CAPLUS, CBNB, CHEMCATS, CHEMLIST, CIN, CSCHEM,
       EMBASE, IFICDB, IFIPAT, IFIUDB, IMSCOSEARCH, MEDLINE, PHAR, PROMT,
       RTECS*, TOXCENTER, USPAT2, USPATFULL
         (*File contains numerically searchable property data)
     Other Sources:
                     EINECS**
         (**Enter CHEMLIST File for up-to-date regulatory information)
DT.CA CAplus document type: Book; Conference; Dissertation; Journal; Patent;
       Report
RL.P
       Roles from patents: ANST (Analytical study); BIOL (Biological study);
       FORM (Formation, nonpreparative); MSC (Miscellaneous); OCCU
       (Occurrence); PREP (Preparation); PROC (Process); PRP (Properties); RACT
       (Reactant or reagent); USES (Uses)
RLD.P Roles for non-specific derivatives from patents: ANST (Analytical
       study); BIOL (Biological study); PREP (Preparation); PROC (Process); PRP
       (Properties); RACT (Reactant or reagent); USES (Uses)
RL.NP Roles from non-patents: ANST (Analytical study); BIOL (Biological
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       (Occurrence); PREP (Preparation); PROC (Process); PRP (Properties); RACT
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RLD.NP Roles for non-specific derivatives from non-patents: ANST (Analytical
       study); BIOL (Biological study); CMBI (Combinatorial study); FORM
       (Formation, nonpreparative); OCCU (Occurrence); PREP (Preparation); PROC
       (Process); PRP (Properties); RACT (Reactant or reagent); USES (Uses)
*** STRUCTURE DIAGRAM IS NOT AVAILABLE ***
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             798 REFERENCES TO NON-SPECIFIC DERIVATIVES IN FILE CA
           11595 REFERENCES IN FILE CAPLUS (1907 TO DATE)
L3
     ANSWER 20 OF 26 REGISTRY COPYRIGHT 2004 ACS on STN
     38234-21-8 REGISTRY
RN
     1-9-Luteinizing hormone-releasing factor (swine), 9-(N-ethyl-L-
     prolinamide) - (9CI) (CA INDEX NAME)
OTHER CA INDEX NAMES:
    Luteinizing hormone-releasing factor (pig), 9-(N-ethyl-L-prolinamide)-10-
     deglycinamide-
OTHER NAMES:
     (des-Gly-NH210, Pro-ethylamide9)-LH-RH
CN
     7: PN: WO0174377 FIGURE: 1 claimed protein
     9-(Ethylamide) Pro-10-des-Gly-NH2-gonadotropin-releasing hormone
CN
     9-(Ethylamide) Pro-10-des-Gly-NH2-LH-releasing factor
CN
CN
     Des-10-glycine-LH-RH-ethylamide
     Des-Gly-10-NH2-LH-RH ethylamide
CN
CN
     Fertirelin
CN
     H 4055
     PGlu-His-Trp-Ser-Tyr-Gly-Leu-Arg-Pro-ethylamide
CN
CN
     [10-Deglycinamide-9-proline ethylamide]-LH-releasing factor
CN
     [10-Des-Gly-NH2, 9-Pro-ethylamide]-LH-releasing factor
CN
     [Des-Gly-NH210, Pro-ethylamide9]-LH-releasing factor
CN
     PROTEIN SEQUENCE; STEREOSEARCH
FS
NTE modified (modifications unspecified)
                 ----- location -----
                                               description
 type
```

uncommon Glp-1 - -

|claimed |FIGURE 1

SEQ 1 XHWSYGLRP

\*\*RELATED SEQUENCES AVAILABLE WITH SEQLINK\*\*

DR 56136-31-3, 70910-43-9

MF C55 H76 N16 O12

CI COM

LC STN Files: AGRICOLA, ANABSTR, BEILSTEIN\*, BIOBUSINESS, BIOSIS, BIOTECHNO, CA, CANCERLIT, CAPLUS, CHEMCATS, DDFU, DRUGU, EMBASE, IFICDB, IFIPAT, IFIUDB, MEDLINE, MRCK\*, MSDS-OHS, TOXCENTER, USAN, USPAT2, USPATFULL, VETU

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DT.CA CAplus document type: Conference; Dissertation; Journal; Patent

RL.P Roles from patents: BIOL (Biological study); PREP (Preparation); PROC (Process); PRP (Properties); USES (Uses)

RLD.P Roles for non-specific derivatives from patents: BIOL (Biological study); USES (Uses)

RL.NP Roles from non-patents: ANST (Analytical study); BIOL (Biological study); PREP (Preparation); PROC (Process); PRP (Properties); RACT (Reactant or reagent); USES (Uses)

RLD.NP Roles for non-specific derivatives from non-patents: BIOL (Biological study)

Absolute stereochemistry.

PAGE 1-B

124 REFERENCES IN FILE CA (1907 TO DATE)

3 REFERENCES TO NON-SPECIFIC DERIVATIVES IN FILE CA

125 REFERENCES IN FILE CAPLUS (1907 TO DATE)

ANSWER 21 OF 26 REGISTRY COPYRIGHT 2004 ACS on STN L3

35110-26-0 REGISTRY RN

D-Glucose, 2-amino-2-deoxy-, homopolymer (9CI) (CA INDEX NAME) CN

OTHER NAMES:

CN Poly(2-deoxy-2-aminoglucose)

Poly(D-glucosamine) CN

Polyglucosamine CN

STEREOSEARCH FS

MF (C6 H13 N O5)x

CI PMS, COM

PCT Polyazomethine, Polyazomethine formed

STN Files: AGRICOLA, BIOBUSINESS, BIOSIS, CA, CAPLUS, CEN, CIN, DIOGENES, IFICDB, IFIPAT, IFIUDB, TOXCENTER, USPAT2, USPATFULL LC

DT.CA CAplus document type: Journal; Patent; Report

Roles from patents: ANST (Analytical study); BIOL (Biological study); RL.P OCCU (Occurrence); PREP (Preparation); PROC (Process); PRP (Properties); RACT (Reactant or reagent); USES (Uses)

RLD.P Roles for non-specific derivatives from patents: ANST (Analytical study); BIOL (Biological study); PREP (Preparation); PRP (Properties); USES (Uses)

Roles from non-patents: ANST (Analytical study); BIOL (Biological study); FORM (Formation, nonpreparative); PREP (Preparation); PROC RL.NP (Process); PRP (Properties); USES (Uses); NORL (No role in record)

RLD.NP Roles for non-specific derivatives from non-patents: PREP (Preparation); USES (Uses)

CM

CRN 3416-24-8 CMF C6 H13 N O5

Absolute stereochemistry. Rotation (+).

- 67 REFERENCES IN FILE CA (1907 TO DATE)
- 13 REFERENCES TO NON-SPECIFIC DERIVATIVES IN FILE CA
- 67 REFERENCES IN FILE CAPLUS (1907 TO DATE)
- ANSWER 22 OF 26 REGISTRY COPYRIGHT 2004 ACS on STN L3

RN 9012-76-4 REGISTRY

Chitosan (8CI, 9CI) (CA INDEX NAME) CN

OTHER NAMES:

CN 100D-VL

CN Amidan

CN BC 10

CN BC 10 (polysaccharide)

```
CN
     Biopolymer L 112
      Chicol
CN
CN
      Chirosan 100
      Chitan, N-acetyl-
CN
CN
      Chitech
CN
      Chitin, N-deacetyl-
      Chitoclear
CN
      Chitoclear 400
CN
CN
      Chitofos
 CN
      Chitolaze
      Chitopearl 3510
CN
      Chitopearl BC 3000
CN
CN
      Chitopearl BCW 2500
      Chitopearl BCW 3000
 CN
      Chitopearl BCW 3500
 CN
      Chitopearl BCW 3505
CN
 CN
      Chitopearl BCW 3507
 CN
      Chitopearl K 20
      Chitosan 10B
 CN
      Chitosan 500
 CN
      Chitosan CLH
 CN
 CN
      Chitosan EL
 CN
      Chitosan F
 CN
      Chitosan FL
 CN
      Chitosan H
 CN
      Chitosan LL
 CN
      Chitosan LL 80
      Chitosan LLWP
 CN
 CN
      Chitosan M
 CN
      Chitosan MP
 CN
      Chitosan PSH
      Chitosan SK 10
 CN
 CN
      Chitosan VL
 CN
      Chitosan WL-M
 CN
      Chitosol
      Chitosom
 CN
 CN
      Crystan LA-S
 CN
      CTA 1 Lactic Acid
      CTA 4
 CN
      DAC 50
 CN
 CN
      DAC 70
 CN
      Daichitosan 100DVL
      Daichitosan DVL
 CN
      Daichitosan L
 CN
      Daichitosan P-VL
 CN
 CN
      Daichitosan VL
 ADDITIONAL NAMES NOT AVAILABLE IN THIS FORMAT - Use FCN, FIDE, or ALL for
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      57285-05-9, 191045-06-4
 MF
      Unspecified
 CI
      PMS, COM, MAN
      Manual registration, Polyother, Polyother only
 PCT
                   ADISNEWS, AGRICOLA, ANABSTR, AQUIRE, BIOBUSINESS, BIOSIS,
      STN Files:
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        PIRA, PROMT, RTECS*, TOXCENTER, TULSA, USAN, USPAT2, USPATFULL, VTB
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      Other Sources: NDSL**, TSCA**, WHO
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 DT.CA Caplus document type: Book; Conference; Dissertation; Journal; Patent;
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        Roles from patents: ANST (Analytical study); BIOL (Biological study); MSC (Miscellaneous); OCCU (Occurrence); PREP (Preparation); PROC
 RL.P
         (Process); PRP (Properties); RACT (Reactant or reagent); USES (Uses);
        NORL (No role in record)
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 RL.NP
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         (Reactant or reagent); USES (Uses); NORL (No role in record)
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         study); BIOL (Biological study); FORM (Formation, nonpreparative); MSC
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(Miscellaneous); OCCU (Occurrence); PREP (Preparation); PROC (Process);
        PRP (Properties); RACT (Reactant or reagent); USES (Uses)
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             2572 REFERENCES TO NON-SPECIFIC DERIVATIVES IN FILE CA
            14679 REFERENCES IN FILE CAPLUS (1907 TO DATE)
     ANSWER 23 OF 26 REGISTRY COPYRIGHT 2004 ACS on STN
L3
RN
     9002-64-6 REGISTRY
     Parathormone (9CI) (CA INDEX NAME)
CN
OTHER NAMES:
CN
     Hormones (animal), parathyroid
     Kakerbin
CN
     Parathormone (1-84)
     Parathyrin
CN
CN
     Parathyroid hormone
CN
     Parathyroidin
CN
     Paroidin
CN
     PTH
     8002-77-5, 9039-27-4
DR
     Unspecified
MF
CI
     PMS, COM, MAN
PCT
     Manual registration
     STN Files: ADISINSIGHT, ADISNEWS, AGRICOLA, ANABSTR, BIOBUSINESS,
LC
       BIOSIS, BIOTECHNO, CA, CABA, CAPLUS, CASREACT, CBNB, CHEMCATS, CHEMLIST,
        CIN, CSCHEM, DDFU, DRUGU, EMBASE, HSDB*, IFICDB, IFIPAT, IFIUDB, IPA,
        MEDLINE, MRCK*, NAPRALERT, PHAR, PROMT, RTECS*, TOXCENTER, USAN, USPAT2,
        USPATFULL
          (*File contains numerically searchable property data)
     Other Sources: NDSL**, TSCA*
          (**Enter CHEMLIST File for up-to-date regulatory information)
DT.CA Caplus document type: Book; Conference; Dissertation; Journal; Patent;
        Report
RL.P
        Roles from patents: ANST (Analytical study); BIOL (Biological study);
        CMBI (Combinatorial study); FORM (Formation, nonpreparative); MSC (Miscellaneous); OCCU (Occurrence); PREP (Preparation); PROC (Process);
        PRP (Properties); RACT (Reactant or reagent); USES (Uses); NORL (No role
        in record)
RLD.P Roles for non-specific derivatives from patents: ANST (Analytical
        study); BIOL (Biological study); PREP (Preparation); PROC (Process); PRP
        (Properties); RACT (Reactant or reagent); USES (Uses)
RL.NP Roles from non-patents: ANST (Analytical study); BIOL (Biological study); FORM (Formation, nonpreparative); MSC (Miscellaneous); OCCU
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*** STRUCTURE DIAGRAM IS NOT AVAILABLE ***
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              320 REFERENCES TO NON-SPECIFIC DERIVATIVES IN FILE CA
            11172 REFERENCES IN FILE CAPLUS (1907 TO DATE)
     ANSWER 24 OF 26 REGISTRY COPYRIGHT 2004 ACS on STN
L3
     123-62-6 REGISTRY
RN
     Propanoic acid, anhydride (9CI) (CA INDEX NAME)
OTHER CA INDEX NAMES:
    Propionic anhydride (6CI, 8CI)
OTHER NAMES:
CN
     Methylacetic anhydride
     Propanoic anhydride
CN
CN
     Propionic acid anhydride
     Propionyl oxide
CN
FS
     3D CONCORD
MF
     C6 H10 O3
CI
     COM
       TN Files: AGRICOLA, ANABSTR, BEILSTEIN*, BIOBUSINESS, BIOSIS, BIOTECHNO, CA, CAOLD, CAPLUS, CASREACT, CEN, CHEMCATS, CHEMINFORMRX,
LC
     STN Files:
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       GMELIN*, HODOC*, HSDB*, IFICDB, IFIPAT, IFIUDB, MEDLINE, MRCK*, MSDS-OHS, NIOSHTIC, PIRA, PROMT, PS, RTECS*, SPECINFO, SYNTHLINE, TOXCENTER, ULIDAT, USPAT2, USPATFULL, VTB
          (*File contains numerically searchable property data)
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DSL\*\*, EINECS\*\*, TSCA\*\* Other Sources:

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- CAplus document type: Conference; Journal; Patent; Report Roles from patents: ANST (Analytical study); BIOL (Biological study); RL.P CMBI (Combinatorial study); FORM (Formation, nonpreparative); MSC (Miscellaneous); OCCU (Occurrence); PREP (Preparation); PROC (Process); PRP (Properties); RACT (Reactant or reagent); USES (Uses); NORL (No role in record)
- Roles for non-specific derivatives from patents: ANST (Analytical study); BIOL (Biological study); OCCU (Occurrence); PREP (Preparation); PROC (Process); PRP (Properties); RACT (Reactant or reagent); USES (Uses)
- Roles from non-patents: ANST (Analytical study); BIOL (Biological study); FORM (Formation, nonpreparative); MSC (Miscellaneous); OCCU RL.NP (Occurrence); PREP (Preparation); PROC (Process); PRP (Properties); RACT (Reactant or reagent); USES (Uses); NORL (No role in record)
- RLD.NP Roles for non-specific derivatives from non-patents: ANST (Analytical study); BIOL (Biological study); FORM (Formation, nonpreparative); PREP (Preparation); PROC (Process); PRP (Properties); RACT (Reactant or reagent); USES (Uses)

## \*\*PROPERTY DATA AVAILABLE IN THE 'PROP' FORMAT\*\*

2808 REFERENCES IN FILE CA (1907 TO DATE)

- 48 REFERENCES TO NON-SPECIFIC DERIVATIVES IN FILE CA
- 2814 REFERENCES IN FILE CAPLUS (1907 TO DATE)
  - 48 REFERENCES IN FILE CAOLD (PRIOR TO 1967)

```
ANSWER 25 OF 26 REGISTRY COPYRIGHT 2004 ACS on STN
1.3
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RN 108-55-4 REGISTRY

2H-Pyran-2,6(3H)-dione, dihydro- (9CI) (CA INDEX NAME)

OTHER CA INDEX NAMES:

Glutaric anhydride (6CI, 7CI, 8CI)

OTHER NAMES:

- Dihydro-2H-pyran-2,6(3H)-dione CN
- Glutaric acid anhydride CN
- CN NSC 16640
- CN Pentanedioic acid anhydride
- Pentanedioic anhydride CN
- CN Pyroqlutaric acid
- FS 3D CONCORD
- MF C5 H6 O3
- CI COM
- LC STN Files: BEILSTEIN\*, BIOBUSINESS, BIOSIS, CA, CANCERLIT, CAOLD, CAPLUS, CASREACT, CHEMCATS, CHEMINFORMRX, CHEMLIST, CSCHEM, DDFU, DETHERM\*, DIPPR\*, DRUGU, HODOC\*, IFICDB, IFIPAT, IFIUDB, MEDLINE, MSDS-OHS, NIOSHTIC, RTECS\*, SPECINFO, SYNTHLINE, TOXCENTER, USPAT2, USPATFULL

(\*File contains numerically searchable property data)

Other Sources: DSL\*\*, EINECS\*\*, TSCA\*\*

- (\*\*Enter CHEMLIST File for up-to-date regulatory information) DT.CA CAplus document type: Conference; Dissertation; Journal; Patent
- Roles from patents: ANST (Analytical study); BIOL (Biological study); CMBI (Combinatorial study); PREP (Preparation); PROC (Process); PRP RL.P (Properties); RACT (Reactant or reagent); USES (Uses); NORL (No role in
- RLD.P Roles for non-specific derivatives from patents: ANST (Analytical study); BIOL (Biological study); PREP (Preparation); PROC (Process); PRP (Properties); RACT (Reactant or reagent); USES (Uses)
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- RLD.NP Roles for non-specific derivatives from non-patents: ANST (Analytical study); BIOL (Biological study); FORM (Formation, nonpreparative); PREP (Preparation); PROC (Process); PRP (Properties); RACT (Reactant or reagent); USES (Uses)

### \*\*PROPERTY DATA AVAILABLE IN THE 'PROP' FORMAT\*\*

1629 REFERENCES IN FILE CA (1907 TO DATE) 116 REFERENCES TO NON-SPECIFIC DERIVATIVES IN FILE CA 1632 REFERENCES IN FILE CAPLUS (1907 TO DATE) 29 REFERENCES IN FILE CAOLD (PRIOR TO 1967)

ANSWER 26 OF 26 REGISTRY COPYRIGHT 2004 ACS on STN 1.3 108-30-5 REGISTRY RN 2,5-Furandione, dihydro- (9CI) (CA INDEX NAME) OTHER CA INDEX NAMES: Succinic anhydride (8CI) CN

OTHER NAMES: 2,5-Diketotetrahydrofuran

Butanedioic anhydride CN CN Dihydro-2,5-furandione

CN NSC 8518

Rikacid SA CN

Succinic acid anhydride CN

Succinyl anhydride CN

CN Succinyl oxide

Tetrahydro-2,5-dioxofuran CN

Tetrahydro-2,5-furandione CN

3D CONCORD FS C4 H4 O3

MF

CI COM

STN Files: AGRICOLA, ANABSTR, AQUIRE, BEILSTEIN\*, BIOBUSINESS, BIOSIS, BIOTECHNO, CA, CANCERLIT, CAOLD, CAPLUS, CASREACT, CBNB, CEN, CHEMCATS, CHEMINFORMRX, CHEMLIST, CIN, CSCHEM, CSNB, DDFU, DETHERM\*, DIPPR\*, DRUGU, EMBASE, ENCOMPLIT, ENCOMPLIT2, ENCOMPPAT, ENCOMPPAT2, GMELIN\*, HODOC\*, HSDB\*, IFICDB, IFIPAT, IFIUDB, MEDLINE, MRCK\*, MSDS-OHS, NIOSHTIC, PIRA, PROMT, PS, RTECS\*, SPECINFO, SYNTHLINE, TOXCENTER, TULSA, USPAT2, USPATFULL, VTB

(\*File contains numerically searchable property data)

Other Sources: DSL\*\*, EINECS\*\*, TSCA\*\* (\*\*Enter CHEMLIST File for up-to-date regulatory information)

DT.CA CAplus document type: Conference; Dissertation; Journal; Patent; Report Roles from patents: ANST (Analytical study); BIOL (Biological study); RL.P CMBI (Combinatorial study); FORM (Formation, nonpreparative); MSC (Miscellaneous); OCCU (Occurrence); PREP (Preparation); PROC (Process); PRP (Properties); RACT (Reactant or reagent); USES (Uses); NORL (No role in record)

Roles for non-specific derivatives from patents: ANST (Analytical study); BIOL (Biological study); CMBI (Combinatorial study); FORM (Formation, nonpreparative); OCCU (Occurrence); PREP (Preparation); PROC (Process); PRP (Properties); RACT (Reactant or reagent); USES (Uses)

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RLD.NP Roles for non-specific derivatives from non-patents: ANST (Analytical study); BIOL (Biological study); FORM (Formation, nonpreparative); PREP (Preparation); PROC (Process); PRP (Properties); RACT (Reactant or reagent); USES (Uses)

## \*\*PROPERTY DATA AVAILABLE IN THE 'PROP' FORMAT\*\*

9401 REFERENCES IN FILE CA (1907 TO DATE)

2849 REFERENCES TO NON-SPECIFIC DERIVATIVES IN FILE CA

9419 REFERENCES IN FILE CAPLUS (1907 TO DATE)

59 REFERENCES IN FILE CAOLD (PRIOR TO 1967)

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MOST RECENT DERWENT UPDATE:
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                                                <200474/DW>
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     1997-042841 [04]
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DNC C2000-120486
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CYC 91
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      AU 2000011045
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      NO 2001001744
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                      A1 20010816 (200147) EN
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                      A1 20020725 (200254)
                      W 20020827 (200271)
      JP 2002527533
                                                        C08B037-08
                      B2 20021112 (200278)
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      US 6479457
                      A1 20030515 (200335)
                                                        C08J003-00
      US 2003092800
US 6794364 B2_20040921_(200462) A61K038-00
ADT WO 2000021567 A1(WO 1999-US23406-19991008; AU 2000011045 A AU
      2000-11045 19991008; NO 2001001744 A WO 1999-US23406 19991008,
   NO 2001-1744 20010406; EP 1123112 A1 EP 1999-954780 19991008, WO 1999-US23406 19991008; US 2002098206 A1 Div ex US 1995-468947
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US 2002-251018 20020920; US 6794364 B2 Div ex US 1995-468947 19950606, CIP of US 1997-929363 19970909, Div ex US 1998-169423 19981009, US 2002-251018

20020920

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          A61K038-04; A61K038-22; A61K047-48; A61P019-10; A61P035-00;
          C07K007-02; C07K011-00; C07K017-10; C08L005-08
    WO 200021567 A UPAB: 20040928
ΔR
     NOVELTY - A copolymer comprising an N-acylated derivative of
     poly(2-amino-2-deoxy-D-glucose, and a composition comprising the polymer and a polypeptide with at least one ionogenic amine, and in which at least
     50 weight% of the polypeptide is ionically bound to the polymer, are useful
     in controlled release polypeptide drug delivery systems.
          DETAILED DESCRIPTION - A copolymer comprising an N-acylated
     derivative of poly(2-amino-2-deoxy-D-glucose), in which 1-50%, by weight,
     of the free amines of the derivative are acylated with a first acyl group
     COE1, and 50-99%, by weight, are acylated with a second acyl group COE2,
     is new. E1 = 3-33C carboxyalkyl, 3-33C carboxyalkenyl, 7-39C
     carboxyarylalkyl or 9-39C carboxyarylalkenyl, E2 = 1-30C alkyl, 2-30C
     alkenyl, 6-37C arylalkyl or 8-37C arylalkenyl, and at least one of the
     free amines is acylated with the first acyl group.
          USE - The composition is used for the controlled drug delivery of
     polypeptides.
          ADVANTAGE - The release of the polypeptide from the composition can
     be varied by e.g. increasing the molecular weight of the polymer to
     decrease the release rate, and increasing the number of carboxylic acid
     groups on the polymer to increase the amount of polypeptide bound to the
     composition, and the amount to be released. Treating the composition with
     soluble salts of di- or polyvalent metals and weak acids, or coating or
     microencapsulating with e.g. an absorbable glycolide copolymer, will alter
     the release rate.
     Dwg.0/0
FS
     CPI
     AB; GI; DCN
FA
     CPI: A03-A00A; A03-C01; A10-E17; A12-V01; B04-C01; B04-C02; B07-A02B
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FILE 'HOME' ENTERED AT 14:04:20 ON 18 NOV 2004

7

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=> b reg
FAMEL REGISTRY ENTERED AT 14:44:21 ON 18 NOV 2004
USE IS SUBJECT TO THE TERMS OF YOUR STN CUSTOMER AGREEMENT.
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```

Property values tagged with IC are from the ZIC/VINITI data file provided by InfoChem.

STRUCTURE FILE UPDATES: 17 NOV 2004 HIGHEST RN 783276-57-3 DICTIONARY FILE UPDATES: 17 NOV 2004 HIGHEST RN 783276-57-3

TSCA INFORMATION NOW CURRENT THROUGH MAY 21, 2004

Please note that search-term pricing does apply when conducting SmartSELECT searches.

Crossover limits have been increased. See HELP CROSSOVER for details.

Experimental and calculated property data are now available. For more information enter HELP PROP at an arrow prompt in the file or refer to the file summary sheet on the web at: http://www.cas.org/ONLINE/DBSS/registryss.html

# es desqide 169tot

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ANSWER 1 OF 2 REGISTRY COPYRIGHT 2004 ACS on STN
L6
    215717-91-2 REGISTRY
RN
    L-Threoninamide, 3-(2-naphthalenyl)-D-alanyl-L-cysteinyl-L-tyrosyl-D-
CN
    tryptophyl-L-lysyl-L-cysteinyl-, cyclic (2.fwdarw.6)-disulfide (9CI) (CA
    INDEX NAME)
FS PROTEIN SEQUENCE; STEREOSEARCH
SOL 7
NTE modified
_____
          ----- location -----
                                       description
type
-15-
terminal mod. Thr-7 - C-terminal amide bridge Cys-2 - Cys-6 disulfide bridge modification Ala-1 - 2-naphthalenyl<2-Naph>
       1 ACYWKCT
         -----
HITS AT: 1-7
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\*\*RELATED SEQUENCES AVAILABLE WITH SEQLINK\*\*

MF C49 H60 N10 O9 S2

SR CA

LC STN Files: CA, CAPLUS, TOXCENTER, USPAT2, USPATFULL

DT.CA CAplus document type: Conference; Patent

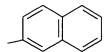
RL.P Roles from patents: BIOL (Biological study); USES (Uses)

RL.NP Roles from non-patents: BIOL (Biological study); PREP (Preparation)

Absolute stereochemistry.

PAGE 1-A s (CH<sub>2</sub>)<sub>4</sub> H<sub>2</sub>N NH2 ö

. PAGE 1-B



CA

SR

L6

2 REFERENCES IN FILE CA (1907 TO DATE) 2 REFERENCES IN FILE CAPLUS (1907 TO DATE)

ANSWER 2 OF 2 REGISTRY COPYRIGHT 2004 ACS on STN  $\cdot$  183580-27-0 REGISTRY RNL-Threoninamide, 3-(1-naphthalenyl)-D-alanyl-L-cysteinyl-L-tyrosyl-D-tryptophyl-L-lysyl-L-cysteinyl-, cyclic (2.fwdarw.6)-disulfide (9CI) (CA INDEX NAME) FS PROTEIN SEQUENCE; STEREOSEARCH SQL NTE modified ----------- location ----description type terminal mod. Thr-7 C-terminal amide disulfide bridge - Cys-6 bridge Cys-2 modification Ala-1 1-naphthalenyl<1-Naph> 1 ACYWKCT SEQ ====== HITS AT: 1-7 \*\*RELATED SEQUENCES AVAILABLE WITH SEQLINK\*\* MF C49 H60 N10 O9 S2

LC STN Files: CA, CAPLUS, TOXCENTER
DT.CA CAplus document type: Conference
RL.NP Roles from non-patents: BIOL (Biological study); PRP (Properties)

Absolute stereochemistry.

PAGE 1-A

PAGE 1-B

PAGE 2-A

PAGE 2-B

1 REFERENCES IN FILE CA (1907 TO DATE)
1 REFERENCES IN FILE CAPLUS (1907 TO DATE)

=> d his

(FILE 'HOME' ENTERED AT 14:00:56 ON 18 NOV 2004)

FILE 'HCAPLUS' ENTERED AT 14:01:54 ON 18 NOV 2004 E WO1999-US23406/APPS E WO99-US23406/APPS

L1 2 E3-4

FILE 'REGISTRY' ENTERED AT 14:02:54 ON 18 NOV 2004

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L2 TRA L1 1- RN : 26 TERMS

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FILE 'WPIX' ENTERED AT 14:02:59 ON 18 NOV 2004 E WO99-US23406/AP, PRN

L4 1 E

L16

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L6 2 L5 AND C6-C6/ES

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                  E RUSSELL RUTH/AU
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                4 E3, E6-7
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=> b hcap FILE 'HCAPLUS' ENTERED AT 14:44:41 ON 18 NOV 2004 USE IS SUBJECT TO THE TERMS OF YOUR STN CUSTOMER AGREEMENT. PLEASE SEE "HELP USAGETERMS" FOR DETAILS. COPYRIGHT (C) 2004 AMERICAN CHEMICAL SOCIETY (ACS)

0 L7 AND L8-16

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FILE COVERS 1907 - 18 Nov 2004 VOL 141 ISS 21 FILE LAST UPDATED: 17 Nov 2004 (20041117/ED)
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This file contains CAS Registry Numbers for easy and accurate substance identification.

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     ANSWER 1 OF 3 HCAPLUS COPYRIGHT 2004 ACS on STN
L7
AN
      1999:690834 HCAPLUS
      131:307099
DN
      Entered STN: 29 Oct 1999
ED
TI
     Use of somatostatin derivatives and/or of phenylhydrazone derivatives as
      antiinflammatory or analgetic agents
     Keri, Gyorgy; Szolcsanyi, Janos; Pinter, Erika; Helyes, Zsuzsanna; Erchegyi, Judit; Horvath, Aniko; Horvath, Judit; Teplan, Istvan; Orfi,
IN
      Laszlo
      Biostatin Gyogyszerkutato-Fejleszto Kft., Hung.
PA
SO
      Eur. Pat. Appl., 20 pp.
      CODEN: EPXXDW
DT
      Patent
      English
LA
      ICM C07K014-655
IC
      ICS A61K038-31; A61K031-15; C07C251-86
      1-11 (Pharmacology)
CC
      Section cross-reference(s): 2
FAN.CNT 1
      PATENT NO.
                                  DATE
                                               APPLICATION NO.
                                                                        DATE
                           KIND
                           ----
                                                                        19990423
ΡI
     EP 952159
                            A2
                                  19991027
                                               EP 1999-107392
      EP 952159
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                                                CA 1999-2269995
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      JP 2000001439
                            A2
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                                                JP 1999-118238
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                                                US 2001-754598
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 PRAI HU 1998-970
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      US 1999-296626
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 EP 952159
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                          C07K014-655
                          A61K038-31; A61K031-15; C07C251-86
                  ICS
 US 2001009899
                  ECLA
                          A61K038/31; C07C251/86; C07K014/655A
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$$Q^{1} \xrightarrow{H} C \xrightarrow{C} N \xrightarrow{N} CH Q^{5}$$

$$Q^{5} \qquad Q^{4}$$

$$Q^{1} \xrightarrow{N} C \xrightarrow{Q^{1}} Q^{2}$$

MARPAT 131:307099

- The invention relates to the use of peptide amides

  R1X1NHCH[(CH2)kR2]COX2X3NHCH[(CH2)nR3]CONHCH[(CH2)kR4]COX4NH2 [X1, X3 = aromatic D-amino acid; X2 = (hydroxyl-substituted) aromatic amino acid; X4 = Thr, Trp; k = 0-3; n = 0, 3, 4; ] and phenylaminooxoacetic acid derivs. I

  (Q1 = H, halo, OH, nitro, amino, C1-4 alkyl, C1-4 alkoxy; Q2 = H, halo, OH, nitro; Q3 = H, halo, OH, nitro, CF3, C1-4 alkyl, C1-4 alkoxy; Q4, Q5 = H, halo, OH, nitro, CF3, C1-4 alkyl, C1-3 dialkylamino), as well as the salts of the above compds., as active substances for the preparation of pharmaceutical compns. possessing neurogenic and non-neurogenic antiinflammatory and analgetic effects.
- ST somatostatin deriv phenylhydrazone deriv analgesic antiinflammatory; phenylaminooxoacetate deriv analgesic antiinflammatory
- IT Analgesics

os

GI

```
Anti-inflammatory agents
         (somatostatin derivs. and/or of phenylhydrazone derivs. as
         antiinflammatory or analgetic agents)
     51110-01-1D, Somatostatin, derivs.
                                           107543-29-3
                                                         147159-51-1, TT-232
IT
                                  169120-33-6
                                                172868-04-1
                                                                215717-90-1
     169120-28-9
                   169120-32-5
      215717-91-2
                   215717-92-3
                                   215717-95-6
                                                 215717-96-7
      247196-17-4
                    247196-18-5
                                   247578-71-8
                                                 247578-72-9
                    247578-75-2
                                   247578-76-3
                                                 247578-77-4D, derivs.
      247578-74-1
                    247578-79-6
                                  247578-80-9
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      247578-78-5
      247591-29-3
      RL: BAC (Biological activity or effector, except adverse); BSU (Biological
      study, unclassified); THU (Therapeutic use); BIOL (Biological study); USES
      (Uses)
         (somatostatin derivs. and/or of phenylhydrazone derivs. as
         antiinflammatory or analgetic agents)
     33507-63-0, Substance P 51110-01-1, Somatostatin 83652-28-2, CGRP
TT
      RL: BPR (Biological process); BSU (Biological study, unclassified); BIOL
      (Biological study); PROC (Process)
         (somatostatin derivs. and/or of phenylhydrazone derivs. as
         antiinflammatory or analgetic agents)
     ANSWER 2 OF 3 HCAPLUS COPYRIGHT 2004 ACS on STN
L7
     1998:597770 HCAPLUS
AN
DN
     130:4050
     Entered STN: 22 Sep 1998
ED
     Somatostatin octa- and heptapeptides, structural and biological
TI
     characteristics
     Horvath, A.; Vadasz, Zs.; Csuka, O.; VanBinst, G.; Jaspers, H.; Idei, I.; Erchegyi, J.; Seprodi, J.; Horvath, J.; Mezo, I.; Teplan, I.; Keri, Gy.
ΑU
     Department of Medical Chemistry, Peptide Biochemistry Research Group,
CS
      Semmelweis University of Medicine, Budapest, H-1444, Hung.
     Peptides 1996, Proceedings of the European Peptide Symposium, 24th,
      Edinburgh, Sept. 8-13, 1996 (1998), Meeting Date 1996, 483-484
      Editor(s): Ramage, Robert; Epton, Roger. Publisher: Mayflower Scientific,
      Kingswinford, UK.
      CODEN: 66RCA5
DT
     Conference
     English
LΑ
CC
      34-3 (Amino Acids, Peptides, and Proteins)
      Section cross-reference(s): 2
      A symposium report on the preparation and in vitro growth hormone inhibitory
AB
      and antiproliferative effects of analogs of H-D-Phe-Cys-Tyr-D-Trp-Lys-Cys-
      Thr-NH2 cyclic disulfide (TT-232).
ST
      somatostatin analog prepn growth hormone inhibitor symposium;
      antiproliferative activity TT 232 analog prepn symposium
TT
      Cytotoxic agents
         (preparation, growth hormone inhibitory activity, and antiproliferative
         activity of somatostatin peptide analogs)
IT
      Growth hormone receptors
      RL: BPR (Biological process); BSU (Biological study, unclassified); BIOL
      (Biological study); PROC (Process)
         (preparation, growth hormone inhibitory activity, and antiproliferative
         activity of somatostatin peptide analogs)
IT
      Proliferation inhibition
         (proliferation inhibitors; preparation, growth hormone inhibitory activity,
         and antiproliferative activity of somatostatin peptide analogs)
      51110-01-1P, SRIF 147159-50-0DP, TT 248, analogs 147159-51-1DP, analogs 183580-29-2P 183580-32-7P 215717-90-1P 215717-91-2P
                                    215717-94-5P
                                                    215717-95-6P
                                                                   215717-96-7P
      215717-92-3P
                     215717-93-4P
      215717-97-8P
      RL: BAC (Biological activity or effector, except adverse); BSU (Biological
      study, unclassified); SPN (Synthetic preparation); BIOL (Biological
      study); PREP (Preparation)
         (preparation, growth hormone inhibitory activity, and antiproliferative
         activity of somatostatin peptide analogs)
RE.CNT
               THERE ARE 2 CITED REFERENCES AVAILABLE FOR THIS RECORD
RE
 (1) Jaspers, H; Int J Peptide Protein Res 1994, V43, P271 HCAPLUS
 (2) Keri, G; Biochem Biophys Res Comm 1993, V191, P681 HCAPLUS
L7
      ANSWER 3 OF 3 HCAPLUS COPYRIGHT 2004 ACS on STN
      1996:639525 HCAPLUS
AN
, DN
      125:317630
      Entered STN: 30 Oct 1996
TI
      Conformationally restricted new somatostatin analogs
     Horvath, A.; Jaspers, H.; Peter, A.; Keri, Gy.; Tourwe, D.; Bokonyi, Gy.;
```

Page 7 Audet 09/870087

- Laus, G.; Csernus, V.; Csuka, O.; et al.
- 1st Institute Biochemistry, Semmelweis Medical University, Budapest, CS H-1444, Hung.
- Peptides 1994, Proceedings of the European Peptide Symposium, 23rd, Braga, SO Port., Sept. 4-10, 1994 (1995), Meeting Date 1994, 564-565. Editor(s): Maia, Hernani L. S. Publisher: ESCOM, Leiden, Neth. CODEN: 63MBAO
- DT Conference
- LA English
- 2-2 (Mammalian Hormones) CC Section cross-reference(s): 1
- The synthesis of structural analogs of somatostatin has led to the design AB of several compds. with improved potencies and/or selective biol. activity. One of these analogs, with a five-residue ring (D-Phe-Cys-Tyr-D-Trp-Lys-Cys-Thr-NH2, TT-232), showed no endocrine but very strong antiproliferative effects in a large variety of cells. Conformational study of the analog revealed a deviation from the typical structural features necessary for somatostatin-like endocrine effects and characteristic to the analogs derived from the Sandoz compound [D-Phe-Cys-Phe-D-Trp-Lys-Thr-Cys-Thr(ol)]. In order to find a general model for somatostatin analogs with selective antitumor activity, the authors synthesized 10 new somatostatin analogs that are related to TT-232 or to the Sandoz compound The authors studied their effect on GH inhibition and cell growth as well as their conformation.
- somatostatin analog conformation activity; TT 232 analog conformation ST activity
- Cell proliferation IT

Conformation and Conformers

Neoplasm inhibitors

(conformationally restricted new somatostatin analogs) 51110-01-1D, Somatostatin, analogs 147159-51-1 183580-27-0 183580-28-1 183580-29-2 183580-30-5 183580-31-6 183580-32-7 183580-33-8

RL: BAC (Biological activity or effector, except adverse); BSU (Biological study, unclassified); PRP (Properties); BIOL (Biological study) (conformationally restricted new somatostatin analogs)

IT 9002-72-6, Growth hormone

RL: BPR (Biological process); BSU (Biological study, unclassified); BIOL (Biological study); PROC (Process)
(conformationally restricted new somatostatin analogs)

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